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(54) **Composition for soluble films with a new hydrolyzed polysaccharide**

(57) The invention is concerned with film-forming compositions containing hydrolysates of the exopolysaccharide YAS34 for the use in pharmaceutical, veterinary, food, cosmetic or other products like films for

wrapping food, aspics or jellies, preferably for predosed formulations like soft or hard capsules.

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joined. The immediate setting of the gelatin solution on the mould pins after dipping is the key step in the process. Otherwise, the gelatin solution would flow down to form capsules with non-uniform wall thickness and unacceptable properties.

[0012] However the aqueous solution of the hydrolyzed polysaccharide YAS34 does not gelatinize at room temperature and does not have the necessary setting ability for hard capsule manufacturing by the described conventional dip moulding process. Surprisingly we have found that the addition of an additional setting agent to the aqueous solution or to the film-forming composition allows the adjustment and optimization of the setting ability of the solution to achieve uniform shell thickness of hard capsules.

[0013] Preferred setting agents for the inventive film-forming compositions or the aqueous solutions thereof are carrageenan, gellan gum, pectin, agarose, gelatin, xanthan with locust bean gum, xanthan with konjac, or unhydrolyzed YAS34. The preferred content of the setting agent in the solid film composition is 0.5 % to 15 % by weight. In the aqueous solution the setting agent is in the range of 0.05 % to 5 % by weight. Optionally the gelling agents are combined with mono or divalent cations like Calcium Ca^{++} or Mg^{++} , e.g. water-soluble salts thereof, preferably in the aqueous solution in an amount of 0.01 to 3 % or in the solid film composition in an amount of 0.1 to 5% by weight.

[0014] Aqueous solutions of the inventive film-forming composition containing hydrolyzed YAS34 and a setting agent and the films derived therefrom have following advantages:

- High solid content which improves significantly the feasibility of different manufacturing processes.
- Sufficient setting ability for hard capsule production by conventional dip moulding process.
- High solubility of the films, suitable for predosed formulations such as hard capsules, soft capsules, tablet coating and ingredient packaging.
- Acceptable mechanical properties of the films for hard capsules allowing filling and blistering operations.

[0015] The inventive composition may contain in a further aspect additional plasticizer, especially for soft film formulations such as soft capsules.

[0016] The inventive composition may contain in a further aspect additional coloring agents or flavoring agents.

[0017] The hard capsules of the invention may be used as containers for providing unit dosage forms, e.g. for agrochemicals, seeds, herbs, foodstuffs, dyestuffs, pharmaceuticals, flavouring agents and the like.

Example 1

Hydrolysis of the native polysaccharide YAS34

[0018] 15 g of YAS34 are dissolved in distilled water at 52°C to form a solution containing 1.5 % by weight to which 1000 ml 2M HCl are added under stirring. The mixture is incubated at 52°C for A 1 hour and in a second batch B for 1.5 hours. After the given incubation time the solution is cooled to room temperature and neutralized to pH 7. The hydrolysates are precipitated with a water/ethanol (50/50 v/v) and washed 4 times with ethanol.

[0019] Molecular weights and viscosity of YAS34 and hydrolysates are shown in Table 1:

Table 1

	Mw average	Viscosity (cps)
YAS34	1,400,000	5% at 50°C gelatinized 5% at 60°C gelatinized
Hydrolysate A	123,000	20% at 50°C 650
Hydrolysate B	45,000	27% at 50°C 300

[0020] The results in Table 1 demonstrate that hydrolysis under mild conditions efficiently reduces molecular weight and viscosity and the hydrolysates are well suitable for the preparation of aqueous solutions with acceptable viscosity and solid content for the manufacturing of hard or soft capsules.

Example 2

Polysaccharide films

[0021]

a) 50 g YAS34 are dissolved in distilled water at 90°C to form a solution of 10 % by weight. After debubbling the solution was stabilized at 80°C. Films with a thickness about 100 µm were prepared by casting the solution on glass plates and drying at 22°C / 50 % RH over night.

b) 50 g hydrolysate B (Mw ≈ 45,000) are dissolved in distilled water at 60°C to form a solution of 27 %. After debubbling, the solution was stabilized at 50°C. Films with a thickness about 100 µm were prepared by casting the solution on glass plates and drying at 22°C / 50 % RH over night.

[0022] The dissolution times of the films shown in Table 2 are determined according to UPS XXII dissolution test method conditions (at 37°C, in demineralized water).

Table 2

	YAS34	Hydrolysate B	Gelatin
Dissolution (min)	51	1.2	2.2

[0023] The mechanical properties of the films shown in Table 3 are determined by tensile tests with an Instron machine with films equilibrated by 50 % RH at 22°C before test.

Table 3

	YAS34	Hydrolysate B
Young's modulus (MPa)	2400	2880
Strength (MPa)	59	55
Elongation at break (%)	19	9

[0024] The result of the tests demonstrate that hydrolysate B has a significantly higher dissolution rate than YAS34. The dissolution is even better than gelatin. The tensile strength of the hydrolysate is reduced when compared with YAS34 but the mechanical properties remain very acceptable not only for soft but even for hard capsule manufacturing.

Example 3

Hard capsules from hydrolysate B

[0025] 54 g hydrolysate B and 4 g of YAS34 are dissolved in distilled water at 70°C to 200 g of a solution containing 27 % hydrolysate B and 2 % YAS34 by weight. After debubbling the solution was stabilized at 60°C.

[0026] The solution thus prepared was then poured into a dipping dish of a single pin pilot machine of conventional hard gelatin capsule production equipment. Hard capsules of size 1 were produced under similar conditions to hard gelatin capsule production. The produced hard capsules have similar dimensions to the conventional gelatin hard capsules.

[0027] The produced hard capsules were evaluated by disintegration tests according to USPXXIII at 37°C in demineralized water:

First leak	1.4 minute
Total disintegration	9.5 minutes

Claims

1. Film-forming composition containing hydrolyzed exopolysaccharide YAS34.

2. Film-forming composition according to claim 1, wherein the EPS YAS34 is hydrolyzed to an average molecular weight range from 500,000 to 10,000.
3. Film composition according to claim 2, wherein the hydrolyzed YAS34 has an average molecular weight of 45,000.
4. Film composition, according to claims 1 to 3 containing an additional setting agent for processing purpose.
5. Film composition according to claim 4, wherein the setting agent is selected from carrageenan, gellan gum, pectin, agorose, gelatin, xanthan with locust bean gum, xanthan with konjac, or native EPS YAS34.
6. Film composition according to claims 4 or 5, wherein the setting agent is contained in an amount of 0.5 to 15 % by weight.
7. Film composition according to claims 4 to 6, wherein the setting agent is combined with salts of mono or divalent cations.
8. Film composition according to claim 7, wherein the cation salt is contained in an amount of 0.01 to 5 % by weight of the whole composition.
9. Film composition according to claims 1 to 8 containing additionally coloring agents, flavoring agents and/or plasticizers.
10. Container produced from the film-forming composition according to claims 1 to 9 for unit dosage forms of agrochemicals, seeds, herbs, foodstuffs, pharmaceuticals or flavoring agents.
11. An aqueous solution of the film-forming composition according to claims 1 to 9 for the manufacturing of soft and hard capsules.
12. The aqueous solution according to claim 11 containing hydrolyzed EPS YAS34 in an amount of 10 to 50%, preferably 15 to 40% by weight of the aqueous solution.
13. The aqueous solution according to claims 11 and 12 containing an additional setting agent in an amount of 0.01 to 5%, preferably 0.03 to 3% by weight of the aqueous solution.
14. The aqueous solution according to claims 11 to 13 containing additionally salts of mono or divalent cations in an amount of 0.01 to 3%, preferably 0.01 to 1% by weight of the aqueous solution.
15. Use of the aqueous solution according to claims 11 and 14 for the manufacturing of hard capsules in a dip moulding process.



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EUROPEAN SEARCH REPORT

Application Number

EP 00 40 2437

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Category	Citation of document with indication, where appropriate, of relevant passages	Relevant to claim	CLASSIFICATION OF THE APPLICATION (Int.Cl.7)
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The present search report has been drawn up for all claims			
Place of search THE HAGUE		Date of completion of the search 5 February 2001	Examiner Lensen, H
<p>CATEGORY OF CITED DOCUMENTS</p> <p>X : particularly relevant if taken alone Y : particularly relevant if combined with another document of the same category A : technological background O : non-written disclosure P : intermediate document</p> <p>T : theory or principle underlying the invention E : earlier patent document, but published on, or after the filing date D : document cited in the application L : document cited for other reasons & : member of the same patent family, corresponding document</p>			

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**ANNEX TO THE EUROPEAN SEARCH REPORT
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